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ATTORNEYS AT LAW

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*LIMITED TO MATTERS
 AND PROCEEDINGS BEFORE
 FEDERAL COURTS & AGENCIES
 **REGISTERED PATENT AGENT
 ***SENIOR COUNSEL

November 9, 2001

WRITER'S DIRECT NUMBER:
 (202) 789-5511

Commissioner for Patents
 Washington, D.C. 20231

Art Unit 1655

Re: U.S. Utility Patent Application
 Appl. No. 09/925,313; Filed: August 10, 2001
 For: **An Alpha-2-Macroglobulin Diagnostic Test**
 Inventors: TANZI *et al.*
 Our Ref: 0609.4460004/JAG/HLK

Sir:

Transmitted herewith for appropriate action are the following documents:

1. Information Disclosure Statement;
2. Form PTO-1449 (citing 98 documents - 29 pages);
3. One copy of each of the 98 cited documents: AA1, AB1, AC1, AD1, AE1, AF1, AL1, AM1, AN1, AO1, AP1, AR1, AS1, AT1, AR2, AS2, AT2, AR3, AS3, AT3, AR4, AS4, AT4, AR5, AS5, AT5, AR6, AS6, AT6, AR7, AS7, AT7, AR8, AS8, AT8, AR9, AS9, AT9, AR10, AS10, AT10, AR11, AS11, AT11, AR12, AS12, AT12, AR13, AS13, AT13, AR14, AS14, AT14, AR15, AS15, AT15, AR16, AS16, AT16, AR17, AS17, AT17, AR18, AS18, AT18, AR19, AS19, AT19, AR20, AS20, AT20, AR21, AS21, AT21, AR22, AS22, AT22, AR23, AS23, AT23, AR24, AS24, AT24, AR25, AS25, AT25, AR26, AS26, AT26, AR27, AS27, AT27, AR28, AS28, AT28, AR29, AS29, AT29; and
4. Return postcard.

Commissioner for Patents
November 9, 2001
Page 2

It is respectfully requested that the attached postcard be stamped with the date of filing of these documents, and that it be returned to our courier. In the event that extensions of time are necessary to prevent abandonment of this patent application, then such extensions of time are hereby petitioned.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036. A duplicate copy of this letter is enclosed.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.



Heidi L. Kraus
Attorney for Applicants
Registration No. 43,730

JAG/HLK/rjv
Enclosures

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

Tanzi *et al.*

Appl. No. 09/925,313

Filed: August 10, 2001

For: **An Alpha-2-Macroglobulin
Diagnostic Test**

Confirmation No.

Art Unit: 1655

Examiner: To be assigned

Atty. Docket: 0609.4460004

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Information Disclosure Statement

Commissioner for Patents
Washington, D.C. 20231

Sir:

Listed on accompanying Form PTO-1449 are documents that may be considered material to the examination of this application, in compliance with the duty of disclosure requirements of 37 C.F.R. §§ 1.56, 1.97 and 1.98. A copy of each of these documents is provided.

Where the publication date of a listed document does not provide a month of publication, the year of publication of the listed document is sufficiently earlier than the effective U.S. filing date and any foreign priority date so that the month of publication is not in issue. Applicants have listed publication dates on the attached PTO-1449 based on information presently available to the undersigned. However, the listed publication dates should not be construed as an admission that the information was actually published on the date indicated.

Applicants reserve the right to establish the patentability of the claimed invention over any of the information provided herewith, and/or to prove that this information may not be prior art, and/or to prove that this information may not be enabling for the teachings purportedly offered.

This statement should not be construed as a representation that a search has been made, or that information more material to the examination of the present patent application does not exist. The Examiner is specifically requested not to rely solely on the material

submitted herewith. It is further understood that the Examiner will consider information that had been cited or submitted to the U.S. Patent and Trademark Office in a prior application relied on under 35 U.S.C. § 120, 1138 OG 37, 38 (May 19, 1992).

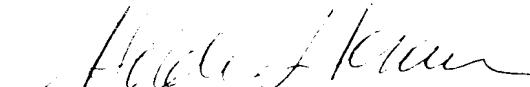
It is respectfully requested that the Examiner initial and return a copy of the enclosed PTO-1449, and to indicate in the official file wrapper of this patent application that the documents have been considered.

This Information Disclosure Statement is being filed within three months of the date of filing of a national application and before the mailing date of a first Office Action on the merits. No statement or fee is required.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.



Heidi L. Kraus
Attorney for Applicants
Registration No. 43,730

Date: October 9, 2001

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FORM PTO-1449

INFORMATION DISCLOSURE STATEMENT

ATTY. DOCKET NO.
0609.4460004

APPLICATION NO.
09/925,313

APPLICANT

Tanzi et al.

FILING DATE
August 10, 2001

GROUP
1655

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U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE
	AA1	4,683,195	07/28/1987	Mullis <i>et al.</i>			
	AB1	5,830,670	11/03/1998	de la Monte <i>et al.</i>			
	AC1	5,948,634	09/07/1999	de la Monte <i>et al.</i>			
	AD1	5,948,888	09/07/1999	de la Monte <i>et al.</i>			
	AE1	6,071,705	06/06/2000	Wands <i>et al.</i>			
	AF1	6,156,311	12/05/2000	Strickland <i>et al.</i>			
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FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
	AL1	WO 97/04794	02/13/1997	WIPO			Yes No
	AM1	WO 91/03557	03/21/1991	WIPO			Yes No
	AN1	WO 92/03474	03/05/1992	WIPO			Yes No
	AO1	WO 96/39834	12/19/1996	WIPO			Yes No
	AP1	WO 99/11824	03/11/1999	WIPO			Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>1</u>	Alvarez, V., et al., "Association between an α -Macroglobulin DNA Polymorphism and Late-Onset Alzheimer's Disease," <i>Journal of Alzheimer's Disease and Parkinsonism</i> 1:48-50, Academic Press, Inc. (October 1999).
	AS	<u>1</u>	Andersen, G.R., et al., "Low Resolution X-Ray Structure of Human Methylamine-treated α -Macroglobulin," <i>Journal of Biochemistry</i> 117:25133-25151, The American Society for Biochemistry and Molecular Biology (1995).
	AT	<u>1</u>	Barrett, A.J., et al., "The Electrophoretically 'Slow' and 'Fast' Forms of the α -Macroglobulin Molecule," <i>Journal of Immunology</i> 131:401-418, Portland Press (1979).

EXAMINER _____ DATE CONSIDERED _____

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant.



FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT		ATTY. DOCKET NO. 0609.4460004	APPLICATION NO. 09/925,313
		APPLICANT Tanzi et al.	
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	2	Bauer, J. et al., "Interleukin-6 and α -2-macroglobulin indicate an acute-phase state in Alzheimer's disease cortices," <i>Neurology</i> 46:111-114 Elsevier Science B.V. (1991).
AS	2	Blacker, D. et al., "Alpha-2 macroglobulin is genetically associated with Alzheimer disease," <i>Nature Genetics</i> 20:357-360, Nature Publishing Group (1998).
AT	2	Blacker, D., et al., "Alzheimer's," <i>Neurology</i> 53:21-22, Nature Publishing Group (May 1999).

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	3	Blennow, K., et al., "No association between the β -macroglobulin (A2M) deletion and Alzheimer's disease, and no change in A2M mRNA, protein, or protein expression," <i>Neurology</i> 55:1065-1079, Springer-Verlag (August 2000).
	AS	3	Boado, R.J., "Antisense drug delivery through the blood-brain barrier," <i>Adv. Brain Delivery Rev.</i> 14:73-107, Elsevier Science (1995).
	AT	3	Boado, R.J., et al., "Drug Delivery of Antisense Molecules to the Brain for Treatment of Alzheimer's Disease and Cerebral AIDS," <i>Adv. Brain Delivery Rev.</i> 14:1308-1315, American Chemical Society and American Pharmaceutical Association (1998)

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	4	Borth, W. " -Macroglobulin a multifunctional binding protein with targeting characteristics," <i>FASEB J.</i> 6:3345-3353, The Federation of American Societies for Experimental Biology (1992).
AS	4	Bowen, M.E., and Gettins, P.G.W. "Bait Region Involvement in the Dimer-Dimer Interface of Human -Macroglobulin and in Mediating Gross Conformational Change," <i>J. Biol. Chem.</i> 273:1825-1831 The American Society for Biochemistry and Molecular Biology, Inc. (1998).
AT	4	Bretaudiere, J.-P., et al. "Structure of native -macroglobulin and its transformation to the protease bound form," <i>Proc. Natl. Acad. Sci. USA</i> 85:1437-1441, The National Academy of Science (1988).

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	5	Businaro, R., et al., "Synthesis and secretion of -macroglobulin by human glioma established cell lines," <i>Exp. Brain Res.</i> 104:213-218, Springer-Verlag (1992).
AS	5	Cavus, I., et al., "Inhibition of Long-term Potentiation Development in Rat Hippocampal Slice by -Macroglobulin, an Acute-Phase Protein in the Brain," <i>J. Neurosci.</i> 16:282-288. Society for Neuroscience (1996).
AT	5	Chen, L., et al., "Apolipoprotein E promoter and -2-Macroglobulin polymorphisms are not genetically associated with Chinese late onset Alzheimer's disease," <i>Neuroscience</i> 87:173-177, Elsevier Science (July 1999).

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	AL						Yes No
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	6	Crawford, F., et al., "The alpha-2 macroglobulin gene is not associated with Alzheimer's disease in a case-control sample," <i>Neurosci. Lett.</i> 270:133-136, Elsevier Science (August 1999).
	AS	6	Dodel, R.C., et al., "α-2 Macroglobulin and the risk of Alzheimer's disease," <i>Neurology</i> 54:438-442, American Academy of Neurology (January 2000).
	AT	6	Dow, D.J., et al., "α-2 macroglobulin polymorphism and Alzheimer disease risk in the UK," <i>Acta Genet. Scand.</i> 50:16-17, Nature Publishing Group (May 1999).

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AL						Yes No
AM						Yes No
AN						Yes No
AO						Yes No
AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	Z	Du, Y., et al., "IgG -Macroglobulin as a β -Amyloid Peptide-Binding Plasma Protein," <i>Neuroscience</i> 74:299-305 Lippincott-Raven Publishing (1997).
AS	Z	Du, Y., et al., "IgG -Macroglobulin Attenuates β -Amyloid Peptide 1-40 Fibril Formation and Associated Neurotoxicity of Cultured Fetal Rat Cortical Neurons," <i>Neuroscience</i> 87:1182-1188, Lippincott-Raven Publishing (1998).
AT	Z	Falker, B., et al., "Short Antisense Oligonucleotide-mediated Inhibition Is Strongly Dependent on Oligo Length and Concentration but Almost Independent of Location of the Target Sequence," <i>Journal of Molecular Biology</i> 241:16187-16194, The American Society for Biochemistry and Molecular Biology (1994).

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FORM PTO-1449

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	AL						Yes No
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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	8	"Freely Associating," <i>Journal of Neurosci</i> 19:1-2, Nature Publishing Group (May 1999).
	AS	8	Feldman, S.R., et al., "Model of γ -Macroglobulin structure and function," <i>Proc. Natl. Acad. Sci. USA</i> 82:5700-5704, The National Academy of Science (1985).
	AT	8	Gauderman, W.J., et al., "Family-Based Association Studies," <i>Human Genetics</i> 103:31-37, Oxford University Press (1999).

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FORM PTO-1449 <u>INFORMATION DISCLOSURE STATEMENT</u>		ATTY. DOCKET NO. 0609.4460004	APPLICATION NO. 09/925,313
		APPLICANT Tanzi <i>et al.</i>	
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	9	Gliemann, J., "Receptors of the Low Density Lipoprotein (LDL) Receptor Family in Man. Multiple Functions of the Large Family Members via Interaction with Complex Ligands," <i>Bind. Chem.</i> 329:951-964, Walter de Gruyter (1998).
AS	9	Hampe, J., <i>et al.</i> , "Mapping Genes for Polygenic Disorders: Considerations for Study Design in the Complex Trait of Inflammatory Bowel Disease," <i>Hum. Hered.</i> 50:91-101, Karger (March-April 2000).
AT	9	Hollenbach, E., <i>et al.</i> , "Confirmation of an association between a polymorphism in exon 3 of the low-density lipoprotein receptor-related protein gene and Alzheimer's disease," <i>Neurology</i> , 51:1905-1907, Lippincott-Raven Publishers (1998).

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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	10	Horvath, S., and Laird, N.M., "A Discordant-Sibship Test for Disequilibrium and Linkage: No Need for Parental Data," <i>American Journal of Human Genetics</i> 59:1886-1897, American Society of Human Genetics (1998).
	AS	10	Hughs, S.R., et al., "IgM-macroglobulin associates with β -amyloid peptide and prevents fibril formation," <i>Proceedings of the National Academy of Sciences</i> 95:3275-3280, The National Academy of Science (1998).
	AT	10	Jinnah, H.A. and Friedmann, T., "Gene Therapy and the brain," <i>British Medical Journal</i> 311:138-148, The British Council (1995).

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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	<u>11</u>	Justus, C.W.E., et al., "Quantification of free α -macroglobulin and α -macroglobulin-protease complexes by a novel ELISA system based on streptococcal α -macroglobulin receptors," <i>Thromb Haemost</i> 63:103-108, Elsevier Science (1990).
AS	<u>11</u>	Kan, C.-C., et al., "Nucleotide sequence of cDNA encoding human α -macroglobulin and assignment of the chromosomal locus," <i>Proc Natl Acad Sci USA</i> 82:2282-2286, National Academy of Science (1985).
AT	<u>11</u>	Kang, D.E., et al., "Genetic association of the low-density lipoprotein receptor-related protein gene (LRP), an apolipoprotein E receptor, with late-onset Alzheimer's disease," <i>Am J Med Genet</i> 3:56-61, Lippincott-Raven Publishers (1997).

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FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT		ATTY. DOCKET NO. 0609.4460004	APPLICATION NO. 09/925,313
		APPLICANT Tanzi	
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	AR	<u>12</u>	Kolodziej, S.J., et al., "Three-Dimensional Structure of the Human Plasmin -Macroglobulin Complex," <i>Science</i> 281:124-133, Academic Press (1998).
	AS	<u>12</u>	Korovaitseva, G.I., et al., "Alpha-2 macroglobulin gene in early- and late-onset Alzheimer Disease," <i>Neurology</i> 53:129-131, Elsevier Science (August 1999).
	AT	<u>12</u>	Koster, M.N., et al., "The α -2-macroglobulin gene in AD," <i>Neurology</i> 55:678-684, American Academy of Neurology (September 2000).

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AR	<u>13</u>	Kounnas, M.Z. et al., "LDL Receptor-Related Protein, a Multifunctional ApoE Receptor, Binds Secreted β -Amyloid Precursor Protein and Mediates Its Degradation," <i>J. Biol. Chem.</i> 270:331-340, Cell Press (1995).
AS	<u>13</u>	Kounnas, M.Z. et al., "The Cellular Internalization and Degradation of Hepatic Lipase Is Mediated by Low Density Lipoprotein Receptor-related Protein and Requires Cell Surface Proteoglycans," <i>J. Biol. Chem.</i> 270:9307-9312, The American Society for Biochemistry and Molecular Biology (1995).
AT	<u>13</u>	Kounnas, M.Z. et al., "LDL Receptor-Related Protein (LPR) Binds Secreted β -Amyloid Precursor Protein and Mediates Its Degradation," <i>Mol. Biol. Cell</i> 6(10 Suppl):327A, The American Society for Cell Biology (1995).

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AR	<u>14</u>	Kovacs, D.M., et al., "Biological Consequences of the Alpha-2-Macroglobulin-2 Allele Associated with Alzheimer's Disease," A Keystone Symposium on Aging: Genetic and Environmental Influences on Lifes Spans, Tamarron, Colorado (February 1999).
AS	<u>14</u>	Kovács, T., et al., "Alpha-2-macroglobulin intronic polymorphism is not associated with autopsy-confirmed late-onset Alzheimer disease," <i>Neurology</i> 53:61-63, Elsevier Science (September 1999).
AT	<u>14</u>	Kovacs, D.M., "2-Macroglobulin in late-onset Alzheimer's disease," <i>Neurology</i> 55:473-479, Elsevier Science Inc. (July 2000).

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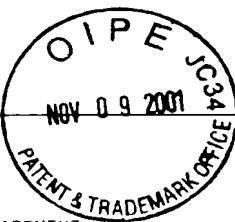
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	AR	15	Krüger, R., et al., "Genetic analysis of the β -2-macroglobulin gene in early- and late-onset Parkinson's disease," <i>Neurology</i> 55:2439-2442, Lippincott Williams & Wilkins (August 2000).
	AS	15	Lake, S.L., et al., "Family-Based Tests of Association in the Presence of Linkage," <i>Am J Hum Genet</i> 67:1515-1525, University of Chicago Press (December 2000).
	AT	15	Le Corre, S.M., et al., "Critical Issues In The Antisense Inhibition of Brain Gene Expression [in vitro]: Experiences Targetting The 5-HT Receptor," <i>Neurochem Int</i> 31:349-362, Elsevier Science Ltd. (1997).

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	AR	16	Liao, A., et al., "Genetic association of an α -2-macroglobulin (Val1000Ile) polymorphism and Alzheimer's disease," <i>Am. J. Med. Genet.</i> 1:1953-1956, Oxford University Press (1998).
	AS	16	Lopes, M.B.S., et al., "Expression of α -macroglobulin receptor/low density lipoprotein receptor-related protein is increased in reactive and neoplastic glial cells," <i>FEBS Letters</i> 344:301-305, Elsevier on behalf of the Federation of European Biochemical Society (1994).
	AT	16	Lorent, K., et al., "Expression In Mouse Embryos And In Adult Mouse Brain of the Three Members of the Amyloid Precursor Protein Family, Of The Alpha-2-Macroglobulin Receptor/Low Density Lipoprotein And Of Its Ligands Apolipoprotein E, Lipoprotein Lipase, Alpha-2-Macroglobulin and the 40,000 Molecular Weight Receptor-Associated Protein," <i>FEBS Letters</i> 367:1009-1025, Elsevier Science Ltd. (1995).

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FORM PTO-146
INFORMATION DISCLOSURE STATEMENT

ATTY. DOCKET NO.
0609.4460004

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	AM						Yes No
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	AO						Yes No
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	AR	17	Marynen, P., et al., "A Monoclonal Antibody To A Neo-Antigen On Macroglobulin Complexes Inhibits Receptor-Mediated Endocytosis," <i>Immuno</i> :1782-1786, American Association of Immunologists (1981).
	AS	17	Marynen, P., et al., "A genetic polymorphism in a functional domain of human pregnancy zone protein: the bait region," <i>Hum Mol Genet</i> :349-352, Elsevier Science B.V. (1990).
	AT	17	Matthijs, G., et al., "A deletion polymorphism in the human alpha-2-macroglobulin (A2M) gene," <i>Am J Hum Genet</i> :5102, Oxford University Press (1991).

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	AR	18	Misra, U.K. and Pizzo, S.V., "Binding of Receptor-recognized Forms of α -Macroglobulin to the α -Macroglobulin Signaling Receptor Activates Phosphatidylinositol 3-Kinase," <i>Journal of Biol. Chem.</i> 273:13399-13402, The American Society for Biochemistry and Molecular Biology (1998).
	AS	18	Mori, T., et al., " α -Macroglobulin is an astroglia-derived neurite-promoting factor for cultured neurons from rat central nervous system," <i>NeuroReport</i> 1:55-61, Elsevier Science B.V. (1990).
	AT	18	Myllykangas, L., et al., "Genetic Association of α -2-Macroglobulin with Alzheimer's Disease in a Finnish Elderly Population," <i>Am. J. Med. Genet.</i> 10:382-390, American Neurological Association (1999).

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AR	19	Narita, M., et al., ", -Macroglobulin Complexes with and Mediates the Endocytosis of -Amyloid Peptide Via Cell Surface Low-Density Lipoprotein Receptor-Related Protein," <i>Journal of Biological Chemistry</i> , v:1904-1911, Lippincott-Raven Publishers (1997).
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AT	19	Polley, W., et al., "Sequence Polymorphism in the Human -2-macroglobulin (A2M) gene," <i>Journal of Biological Chemistry</i> , v:198, Oxford University Press (1991).

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	AR	20	Pollier, W., et al. "Cloning of the human α -macroglobulin gene and detection of mutations in two functional domains: the bait region and the thiolester site," <i>Imm. Geriat.</i> 26:313-319, Springer-Verlag (1992).
	AS	20	Qui, W.Q., et al. "Degradation of Amyloid β -Protein by a Serine Protease- α -Macroglobulin Complex," <i>J. Biol. Chem.</i> 271:8443-8451, The American Society for Biochemistry and Molecular Biology, Inc. (1996).
	AT	20	Rebeck, G.W., et al. "Multiple, Diverse Senile Plaque-associated Proteins Are Ligands of an Apolipoprotein E Receptor, the α -Macroglobulin Receptor/Low-Density-Lipoprotein Receptor-related Protein," <i>Ann. Neurol.</i> 38:211-217, American Neurological Association (1995)

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AR	<u>21</u>	Reeck, G.R., et al., "Homology' in Proteins and Nucleic Acids: A Terminology Muddle and a Way out of It," <i>Cell</i> 50:667, Cell Press (1987).
AS	<u>21</u>	Rogaeva, E.A., et al., "An α -2-macroglobulin insertion-deletion polymorphism in Alzheimer's disease," <i>Nature Genetics</i> 21:19-21, Nature Publishing Group (May 1999).
AT	<u>21</u>	Romas, S.N., et al., "The deletion polymorphism and Val1000Ile in α -2-macroglobulin and Alzheimer disease in Caribbean Hispanics," <i>Journal of Internal Medicine</i> 248:133-136, Elsevier Science (February 2000).

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	AR	22	Rudrasingham, V., et al., " β -2 macroglobulin gene and Alzheimer disease," <i>Am. J. Med. Genet.</i> 11:17-19, Nature Publishing Group (May 1999).
	AS	22	Selkoe, D.J., "Cell biology Of The Amyloid β -Protein Precursor And The Mechanism Of Alzheimer's Disease," <i>Am. J. Med. Genet.</i> 11:373-403, Annual Reviews (1994)
	AT	22	Singleton, A.B., et al., " β -Macroglobulin polymorphisms in Alzheimer's disease and dementia with Lewy bodies," <i>Am. J. Med. Genet.</i> 11:1507-1510, Lippincott Williams & Wilkins (May 1999).

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	AR	<u>23</u>	Sodeyama, N., et al., "x2-Macroglobulin Polymorphism is not associated with AD or AD-type neuropathology in the Japanese," <i>Neurology</i> 64:443-446, American Academy of Neurology (January 2000).
	AS	<u>23</u>	Soto, C., et al., "x-Sheet breaker peptides inhibit fibrillogenesis in a rat brain model of amyloidosis: Implications for Alzheimer's therapy," <i>Nat. Med.</i> 4:822-826, Nature Publishing Group (1998).
	AT	<u>23</u>	Soto, C., et al., "Inhibition of Alzheimer's Amyloidosis by Peptides That Prevent x-Sheet Conformation," <i>Proc. Natl. Acad. Sci. USA</i> 93:672-680, Academic Press (1996).

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	AR	<u>24</u>	Sottrup-Jensen, L., et al., "Primary Structure of Human γ -Macroglobulin," <i>J. Biol. Chem.</i> 259:8318-8327, The American Society of Biological Chemistry (1984).
	AS	<u>24</u>	Stratagene, 1998 Catalog, p. 39, "Gene Characterization Kits," La Jolla, CA (1997).
	AT	<u>24</u>	Strauss, S., et al., "Detection of Interleukin-6 and γ -Macroglobulin Immunoreactivity in Cortex and Hippocampus of Alzheimer's Disease Patients," <i>Am. J. Pathol.</i> 141:223-230 The United States and Canadian Academy of Pathology, Inc. (1992).

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	AP						Yes No

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	AR	<u>25</u>	Strickland, D.K., et al., "Sequence Identity between the α -Macroglobulin Receptor and Low Density Lipoprotein Receptor-related Protein Suggests That This Molecule Is a Multifunctional Receptor," <i>J. Biol. Chem.</i> 266:17401-17404, The American Society for Biochemistry and Molecular Biology (1990).
	AS	<u>25</u>	Strickland, D.K., et al., "Primary Structure of α -Macroglobulin Receptor-associated Protein," <i>J. Biol. Chem.</i> 266:13364-13369, The American Society for Biochemistry and Molecular Biology (1991).
	AT	<u>25</u>	Tanzi, R., et al., "Amyloid β Protein Gene: cDNA, mRNA Distribution, and Genetic Linkage Near the Alzheimer Locus," <i>Science</i> 235:880-884, American Association for the Advancement of Science (1987).

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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
AL							Yes No
AM							Yes No
AN							Yes No
AO							Yes No
AP							Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	26	Tanzi, R.E., "A genetic dichotomy model for the inheritance of Alzheimer's disease and common age-related disorders," <i>Journal of Clinical Investigation</i> 101:1175-1179, American Society for Clinical Investigation (November 1999).
	AS	26	Tirode, F., et al., "A Conditionally Expressed Third Partner Stabilizes or Prevents the Formation of a Transcriptional Activator in a Three-hybrid System," <i>Journal of Molecular Biology</i> 22995-22999 The American Society for Biochemistry and Molecular Biology (1997).
	AT	26	Tooyama, I., et al., "Immunohistochemical Study of a Macroglobulin Receptor in Alzheimer and Control Postmortem Human Brain," <i>Journal of Neuropathology and Experimental Neurology</i> 52:153-160. Humana Press, Inc (1993).

EXAMINER	DATE CONSIDERED
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EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant.



FORM PTO-1449 PATENT & TRADEMARK OFFICE INFORMATION DISCLOSURE STATEMENT		ATTY. DOCKET NO. 0609.4460004	APPLICATION NO. 09/925,313
		APPLICANT Tanzi, et al.	
		FILING DATE August 10, 2001	GROUP 1655

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AN							Yes No
AO							Yes No
AP							Yes No

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	AR	<u>27</u>	Van Gool, D., et al., "γ-Macroglobulin Expression in Neuritic-Type Plaques in Patients With Alzheimer's Disease," <i>Neurology</i> 47: 233-237 Pergamon Press Ltd. (1993).
	AS	<u>27</u>	Van Leuven, F., et al., "The Receptor-binding Domain of Human γ-Macroglobulin. Isolation After Limited Proteolysis with a Bacterial Proteinase," <i>Journal of Biological Chemistry</i> 261:11369-11373, The American Society of Biological Chemists, Inc. (1986).
	AT	<u>27</u>	Verpillat, P., et al., "γ2-Macroglobulin Gene and Alzheimer's Disease: Confirmation of Association by Haplotypes Analyses," <i>Annals of Neurology</i> 48:400-402, American Neurological Association and the Child Neurology Society (September 2000).

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FORM PTO-1449

INFORMATION DISCLOSURE STATEMENT

ATTY. DOCKET NO.
0609.4460004APPLICATION NO.
09/925,313APPLICANT
Tanzi et al.FILING DATE
August 10, 2001GROUP
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U.S. PATENT DOCUMENTS

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	AN						Yes No
	AO						Yes No
	AP						Yes No

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	AR	28	Wavrant-DeVrieze, F., et al., "Association between the low density lipoprotein receptor-related protein (LRP) and Alzheimer's disease," <i>Neurology</i> , v. 51, p. 68-70, Elsevier Science Ireland Ltd. (1997).
	AS	28	Wavrant-DeVrieze, F., et al., "No association between the alpha-2 macroglobulin in I1000V polymorphism and Alzheimer's disease," <i>Neurology</i> , v. 53, p. 137-139, Elsevier Science Ireland Ltd. (March 1999).
	AT	28	Zhang, Z., et al., "Inhibition of α -macroglobulin/proteinase-mediated degradation of amyloid peptide by apolipoprotein E and α -antichymotrypsin," <i>Neuroscience</i> , v. 73, p. 156-161, Parthenon Publishing Group (1996).

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AO						Yes No
AP						Yes No

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AR	<u>29</u>	Zill, P., et al., "Polymorphisms in the -2 macroglobulin gene in psychogeriatric patients," <i>Psychogeriatrics</i> 2000; 41:69-72, Elsevier Science Ireland Ltd. (November 2000).
AS	<u>29</u>	International Search Report for International Application No. PCT/US00/02412, mailed July 6, 2000.
AT	<u>29</u>	

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